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Date of Signature and Deposit: May 21, 2002

Zhibin Ren

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Richard J. Roman
David R. Harder
Noriyuki Miyata
Masakazu Sato
Kazuya Kameo
Shigeru Okuyama

Date: May 21, 2002

Serial No.: 09/937,946

Group Art Unit:

Filed: 10/02/2001

Examiner:

Title: USE OF 20-HETE SYNTHESIZING ENZYME
INHIBITORS AS THERAPY FOR CEREBRAL
VASCULAR DISEASE

File No.: 650053.91533

INFORMATION DISCLOSURE STATEMENT

Commissioner For Patents
Box Non-Fee
P O Box 2327
Arlington, VA 22202

Dear Sir:

Enclosed is a completed Form PTO-1449 listing documents that the applicants in the above-identified application wish to bring to the attention of the Examiner for consideration in connection with the examination on the merits of this application.

No fee is believed due in connection with this submission. However, if a fee is due, please charge the fee to Deposit Account No. 17-0055.

Respectfully submitted,

Zhibin Ren
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Form PTO-1449 (Rev. 2-88)				U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTY. DOCKET NO. 650051 533	SERIAL NO. 09/937,946
INFORMATION DISCLOSURE STATEMENT BY APPLICANT <i>(Use several sheets if necessary)</i>				APPLICANT(S): Roman/Harder/Miyata/Sato/Dameo/Okuyama			
<i>MAY 24 2002</i> <i>OIF</i> <i>CIAA</i> <i>PATENT & TRADEMARK OFFICE</i>				FILING DATE: 10/02/2001		GROUP	
U.S. PATENT DOCUMENTS							
EXAMINER'S INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
FOREIGN PATENT DOCUMENTS							
		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	TRANSLATION
		WO 01/32164	10.05.2001	Japan			YES NO
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)							
		Alonso-Galicia, M. et al., "Inhibition of 20-HETE Production Contributes to the Vascular Responses to Nitric Oxide," <u>Hypertension</u> 29:320-325 (1997)					
		Alonso-Galicia, M., et al., "Contribution of 20-HETE to Vasodilator Actions of Nitric Oxide in the Cerebral Microcirculation," <u>Stroke</u> 30:2727-2734 (1999)					
		Alonso-Galicia, M., et al., "20-HETE agonists and antagonists in the renal circulation," <u>Am. J. Physiol.</u> 277:F790-F796 (1999)					
		Amet, Y., et al., "Cytochrome P450 4A and 2E1 Expression in Human Kidney Microsomes," <u>Biochemical Pharmacology</u> 53:765-771 (1997)					
		Amet, Y., et al., "Noninvolvement of CYP2E1 in the (ω -1)-hydroxylation of Fatty Acids in Rat Kidney Microsomes," <u>Biochemical Pharmacology</u> 54:947-952 (1997)					
		Amet, Y., et al., "P-450-Dependent Metabolism of Lauric Acid in Alcoholic Liver Disease: Comparison Between Rat Liver and Kidney Microsomes," <u>Alcoholism: Clinical and Experimental Research</u> 22:455-462 (1998)					
		Bednar, M.M., et al., "16(R)-Hydroxy-5,8,11,14-eicosatetraenoic Acid, a New Arachidonate Metabolite in Human Polymorphonuclear Leukocytes," <u>Biochemical Pharmacology</u> 60:447-455 (2000)					
		Christmas, P., et al., "Alternative Splicing Determines the Function of CYP4F3 by Switching Substrate Specificity," <u>Journal of Biological Chemistry</u> 276:38166-38172 (2001)					
		Clozel, M. et al., "BQ-123, A Peptidic Endothelin ET _A Receptor Antagonist, Prevents the Early Cerebral Vasospasm Following Subarachnoid Hemorrhage After Intracisternal But Not Intravenous Injection," <u>Life Sciences</u> 52:825-834 (1993)					
		Croft, K., et al., "Angiotensin II releases 20-HETE from rat renal microvessels," <u>Am. J. Physiol. Renal Physiol.</u> 279:F544-F551 (2000)					
		Dahly, A.J., et al., "Chronic Anti-TGF- β Therapy Improves Blood Pressure and Renal Function in Dahl S Rats," <u>FASEB J.</u> 14:A133 (2000)					
		Dahly, A.J., et al. "Chronic Anti-TGF- β Ab Therapy Reduces Proteinuria and Renal Injury in Hypertensive Rats," <u>American Society of Nephrology Renal Week</u> 11:332A (2000)					
		Gebremedhin, D., et al., "Cat cerebral arterial smooth muscle cells express cytochrome P450 4A2 enzyme and produce the vasoconstrictor 20-HETE which enhances L-type Ca ²⁺ current," <u>Journal of Physiology</u> 507.3:771-781 (1998)					
EXAMINER				DATE CONSIDERED			
* EXAMINER: Initial if a citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. QBMA/332105							

Form PTO-1449 (Rev. 2-88)		U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTY. DOCKET NO. 65005-1533	SERIAL NO. 09/937,946		
INFORMATION DISCLOSURE STATEMENT BY APPLICANT <i>(Use several sheets if necessary)</i>				APPLICANT IS: Roman/Harder/Miyata/Sato/Dameo/Okuyama			
MAY 24 2002				FILING DATE: 10/02/2001	GROUP		
U.S. PATENT DOCUMENTS							
* EXAMINER'S INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
FOREIGN PATENT DOCUMENTS							
		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	TRANSLATION
							YES NO
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)							
		Gebremedhin, D., et al., "Production of 20-HETE and Its Role in Autoregulation of Cerebral Blood Flow," <u>Circulation Research</u> 87:60-65 (2000)					
		Harder, D.R., et al., "Formation and action of a P-450 4A metabolite of arachidonic acid in cat cerebral microvessels," <u>American Physiological Society</u> 266:H2098-H2107 (1994)					
		Juvela S., "Plasma endothelin concentrations after aneurysmal subarachnoid hemorrhage," <u>J. Neurosurg.</u> 92:390-400 (2000)					
		Oyekan, A., et al., "Renal oxygenases: differential contribution to vasoconstriction induced by ET-1 and ANG II," <u>American Physiological Society</u> 273:R293-R300 (1997)					
		Powell, P.K., et al., "Metabolism of Arachidonic Acid to 20-Hydroxy-5,8,11,14-eicosatetraenoic Acid by P450 Enzymes in Human Liver: Involvement of CYP4F2 and CYP4A11," <u>Journal of Pharmacology and Experimental Therapeutics</u> 285:1327-1336 (1998)					
		Rosolowsky, M., "Metabolism of arachidonic acid by canine polymorphonuclear leukocytes synthesis of lipoxygenase and omega-oxidized metabolites," <u>Biochimica et Biophysica Acta</u> 1300:143-150 (1996)					
		Seifert, V., et al., "Endothelin concentrations in patients with aneurysmal subarachnoid hemorrhage," <u>J. Neurosurg.</u> 82:55-62 (1995)					
		Wang, M., et al., "Contribution of cytochrome P-450 4A1 and 4A2 to vascular 20-hydroxyeicosatetraenoic acid synthesis in rat kidneys," <u>American Physiological Society</u> 276L:F246-F253 (1999)					
		Wang, M., et al., "Cytochrome P450-Derived Arachidonic Acid Metabolism in the Rat Kidney: Characterization of Selective Inhibitors," <u>The Journal of Pharmacology and Experimental Therapeutics</u> 284:966-973 (1998)					
		Zou, A., et al., "Effects of 17-Octadecynoic Acid, a Suicide-Substrate Inhibitor of Cytochrome P450 Fatty Acid ω -Hydroxylase, on Renal Function in Rats," <u>The Journal of Pharmacology and Experimental Therapeutics</u> 268:474-481 (1994)					
EXAMINER			DATE CONSIDERED				
<p>* EXAMINER: Initial if a citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. QBMAD\332105</p>							

FEE TRANSMITTAL FOR FY 2002

MAY 24 2002

Patent fees are subject to annual revision.

TOTAL AMOUNT OF PAYMENT

\$

Complete if Known

Application Number	09/937,946
Filing Date	10/02/2001
First Named Inventor	Richard J. Roman
Group Art Unit	
Examiner Name	
Attorney Docket Number	650053.91533

METHOD OF PAYMENT (check one)

1. The Commissioner is hereby authorized to charge indicated fees and credit any over payments to:

Deposit Account Number Deposit Account Name Charge Any Additional Fee Required Under 37 CFR 1.16 and 1.17 Applicant claims small entity status See 37 CFR 1.27

2. Payment Enclosed:

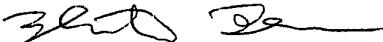
 Check Credit Card Money Order Other
FEE CALCULATION (continued)**3. ADDITIONAL FEES**

Large Entity Fee Code	Small Entity Fee Code	Fee (\$)	Fee (\$)	Fee Description	Fee Paid
105	130	205	65	Surcharge - late filing fee or oath	
127	50	227	25	Surcharge - late provisional filing fee or cover sheet	
139	130	139	130	Non-English specification	
147	2,520	147	2,520	For filing a request for reexamination	
112	920	112	920	Requesting publication of SIR prior to Examiner action	
113	1,840	113	1,840	Requesting publication of SIR after Examiner action	
115	110	215	55	Extension for reply within first month	
116	400	216	200	Extension for reply within second month	
117	920	217	460	Extension for reply within third month	
118	1,440	218	720	Extension for reply within fourth month	
128	1,960	228	980	Extension for reply within fifth month	
119	320	219	160	Notice of Appeal	
120	320	220	160	Filing a brief in support of an appeal	
121	280	221	140	Request for oral hearing	
138	1,510	138	1,510	Petition to institute a public use proceeding	
140	110	240	55	Petition to revive unavoidably abandoned application	
141	1,280	241	640	Petition to revive unintentionally abandoned application	
142	1,280	242	640	Utility issue fee (or reissue)	
143	460	243	230	Design issue fee	
144	620	244	310	Plant issue fee	
122	130	122	130	Petitions to the Commissioner	
123	50	123	50	Petitions related to provisional applications	
126	180	126	180	Submission of Information Disclosure Stmt	
581	40	581	40	Recording each patent assignment per property (times number of properties)	
146	740	246	370	Filing a submission after final rejection (37 CFR 1.129(a))	
146	740	246	370	For each additional invention to be examined (37 CFR 1.129(b))	
179	740	270	370	Request for Continued Examination (RCE)	
169	900	169	900	Request for expedited examination of a design application	
Other fee (specify) _____					

Reduced by Basic Filing Fee Paid

SUBTOTAL (3) (\$).00

SUBMITTED BY**Complete (if applicable)**

Typed or Printed Name	Zhibin Ren	Registration No. (Attorney/Agent)	47,897	Telephone	608/251-5000
Signature			Date	May 21, 2002	